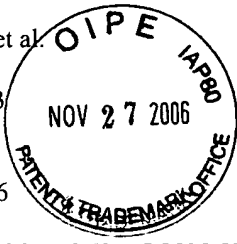


**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

*IFw* ✓

Applicant(s): HAMILL et al. )  
 Serial No.: 10/585,503 )  
 Confirmation No.: 2236 )  
 Filed: 7 July 2006 )  
 For: MECHANOSENSITIVE ION CHANNELS AND METHODS OF USE

Group Art Unit: 1642  
 Examiner: Unassigned  
 Docket No. 265.00450101



**Mail Stop Amendment**

Commissioner for Patents  
 P.O. Box 1450  
 Alexandria, VA 22313-1450

We are transmitting the following documents along with this Transmittal Sheet (which is submitted in triplicate):

- ☒ **Small entity status is entitled to be asserted in the above-identified application.**  
☒ An itemized return postcard.  
☒ An Information Disclosure Statement (2 pgs); copies of 0 applications; 1449 forms (12 pgs); and copies of 108 documents cited on the 1449 forms.  
 Other: \_\_\_\_\_  
 Amendment \_\_\_\_\_ No Additional fee is required. \_\_\_\_\_ The fee has been calculated as shown:

Fee Calculation for Claims Pending After Amendment					
	Pending Claims after Amendment (1)	Claims Paid for Earlier (2)	Number of Additional Claims (1-2)	Cost per Additional Claim	Additional Fees Required
Total Claims				x \$25 =	
Independent Claims				x \$100 =	
One or More New Multiple Dependent Claims Presented? If Yes, Add \$180 Here →					
Total Additional Claim Fees Required					

Please consider this a PETITION FOR EXTENSION OF TIME for a sufficient number of months to enter these papers and please charge any additional fees or credit overpayment to Deposit Account No. 13-4895. Triplicate copies of this sheet are enclosed.

**CERTIFICATE UNDER 37 C.F.R. §1.8:** The undersigned hereby certifies that this Transmittal Letter and the paper(s), as described hereinabove, are being deposited in the United States Postal Service, as first class mail, in an envelope addressed to: Mail Stop Amendment, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on this 22 day of November, 2006.

MUETING, RAASCH & GEBHARDT, P.A.  
 Customer Number: 26813

By: David L. Provence  
 Name: David L. Provence  
 Reg. No.: 43,022  
 Direct Dial: 612-305-1005  
 Facsimile: 612-305-1228

(SMALL ENTITY TRANSMITTAL UNDER RULE 1.8)



PATENT  
Docket No. 265.00450101

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant(s):	HAMILL et al.	)	Group Art Unit:	1642
		)		
Serial No.:	10/585,503	)	Examiner:	Unassigned
Confirmation No.:	2236	)		
		)		
Filed:	7 July 2006	)		
For:	<u>MECHANOSENSITIVE ION CHANNELS AND METHODS OF USE</u>			

**INFORMATION DISCLOSURE STATEMENT**

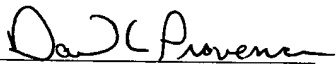
Mail Stop Amendment  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Sir:

In compliance with the duty imposed by 37 C.F.R. § 1.56, and in accordance with C.F.R. §§ 1.97 *et. seq.*, the materials enclosed herewith are brought to the attention of the Examiner as possibly being of interest in connection with the above-identified patent application. Pursuant to MPEP § 609, the information cited in the present Information Disclosure Statement shall not be construed to be an admission that the information is, or is considered to be, material to patentability. Consideration of each of the documents listed on the attached 1449 form(s) is respectfully requested. Pursuant to the provisions of M.P.E.P. §609, Applicants further request that a copy of the 1449 form(s), marked as being considered and initialed by the Examiner, be returned with the next Official Communication.

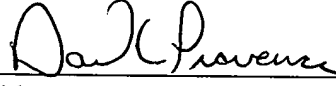
It is believed that no fee is due, as this Information Disclosure Statement is filed prior to the receipt of any Action on the merits. However, in the event a fee is due, please charge any fee or credit any overpayment to Account No. 13-4895.

The Examiner is invited to contact Applicants' Representatives at the below-listed telephone number, if they can be of any assistance during prosecution of the present application.

<b>CERTIFICATE UNDER 37 C.F.R. 1.8:</b>
The undersigned hereby certifies that this paper is being deposited in the United States Postal Service, as first class mail, in an envelope addressed to: Mail Stop Amendment, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on this <u>22</u> day of <u>November</u> , 2006.

Name: David L. Provence

Respectfully submitted  
By  
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November 22, 2006  
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<b>INFORMATION DISCLOSURE STATEMENT</b>	Atty. Docket No.: 265.00450101	Serial No.: 10/585,503
	Applicant(s): HAMILL et al.	Confirmation No.: 2236
	Application Filing Date: July 7, 2006	Group: 1642
	Information Disclosure Statement mailed: November 22, 2006	

**U.S. PATENT DOCUMENTS**

Examiner Initial	Copy Enclosed	Document Number	Date	Name	Class	Subclass	Filing Date If Appropriate
		5,756,663	05/26/98	Lampe et al.			
		6,194,389	02/27/01	Johnston et al.			
		US 2002/0077286 A1	06/20/02	Sachs et al.			
		60/535,327	01/09/04	Hamill et al.			

**FOREIGN PATENT DOCUMENTS**

Examiner Initial	Copy Enclosed	Document Number	Date	Country	Class	Subclass	Translation	
							Yes	No
	X	WO 2005/070122 A2	08/04/05	PCT				

**OTHER DOCUMENTS (Including Authors, Title, Date, Pertinent Papers, etc.)**

Examiner Initial	Copy Enclosed	Document Description
	X	American Type Culture Collection, "ATTC Number CRL-1435," organism: Homo sapiens (humans); designation: PC-3 [online]; Manassas, VA [retrieved on 2006-11-16] from the Internet. Retrieved from the Internet: <URL:http://www.atcc.org/common/catalog/numSearch/numResults.cfm>; 4 pgs.
	X	American Type Culture Collection, "ATTC Number CRL-1740," organism: Homo sapiens (humans); designation: LNCaP clone FGC [online]; Manassas, VA [retrieved on 2006-11-16] from the Internet. Retrieved from the Internet: <URL:http://www.atcc.org/common/catalog/numSearch/numResults.cfm>; 4 pgs.

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<small>*Examiner: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.</small>	

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	X	Auerbach A., "Single-channel dose-response studies in single, cell-attached patches," <i>Biophys. Journal</i> , September 1991;60:660-670.
	X	Banyard & Zetter, "The role of cell motility in prostate cancer," <i>Cancer and Metastasis Reviews</i> , 1999;17:449-458.
	X	BBC News, "Tarantula 'may save lives'," [online]. BBC Homepage World Service Education, 18 May 2000, [retrieved on 2006-November-02]. Retrieved from the Internet:<URL:http://news.bbc.co.uk/2/hi/science/nature/753403.stm>; 3 pgs.
	X	Bielfeld-Ackermann et al., "Maitotoxin (MTX) activates a nonselective cation channel in <i>Xenopus laevis</i> oocytes," <i>Pfluegers Arch. European J. Physiol</i> , 1998;436:329-337.
	X	"BLAST," National Institutes of Health, Bethesda, MD [online]. Retrieved from Internet on April 17, 2001. <URL:http://www.ncbi.nlm.nih.gov/gorf/bl2.html>, 2 pgs.
	X	Bobanovic et al., "Molecular cloning and immunolocalization of a novel vertebrate trp homologue from <i>Xenopus</i> ," <i>Biochem J</i> , 1999;340:593-599.
	X	Bode et al., "Tarantula peptide inhibits atrial fibrillation," <i>Nature</i> , 4 January 2001;409:35-36.
	X	Bormann et al., "Mechanism of anion permeation through channels gated by glycine and $\gamma$ -aminobutyric acid in mouse spinal neurones," <i>J Physiol.</i> , 1987;385:243-286.
	X	Brereton et al., "Maitotoxin activates an endogenous non-selective cation channel and is an effective initiator of the activation of the heterologously expressed hTRPC-1 (transient receptor potential) non-selective cation channel in H4-IIIE liver cells," <i>Biochem. Biophys. Acta.</i> , 2001;1540:107-126.

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	X	Brereton et al., "Evidence that the TRP-1 protein is unlikely to account for store-operated $Ca^{2+}$ inflow in <i>Xenopus laevis</i> oocytes," <i>Mol. Cell. Biochem.</i> , 2000;214:63-74.
	X	Caldwell et al., "Using gadolinium to identify stretch-activated channels: technical considerations," <i>Am J Physiol.</i> , 1998;275:C619-621.
	X	Christensen, "Mediation of cell volume regulation by $Ca^{2+}$ influx through stretch-activated channels," <i>Nature</i> , 5 November 1987;330:66-68.
	X	Clapham, "TRP channels as cellular sensors," <i>Nature</i> , 4 December 2003;426:517-524.
	X	Collins et al., "Identification and isolation of human prostate epithelial stem cells based on $\alpha_2\beta_1$ -integrin expression," <i>J Cell Sci</i> , 2001;114(21):3865-3872.
	X	Diss et al., "Expression of skeletal muscle-type voltage gated $Na^+$ channel in rat and human prostate cancer cell lines," <i>FEBS Letts.</i> , 1998;427:5-10.
	X	Doyle et al., "Calcium transients induce spatially coordinated increases in traction force during the movement of fish keratocytes," <i>J Cell Science</i> , 2004;117:2203-2214.
	X	Fischer et al., eds., <i>The Cancer Chemotherapy Handbook</i> , 6 <sup>th</sup> Ed., 2003, title page, publisher's page and table of contents only; 6 pgs.
	X	Franco-Obregón and Lansman, "Changes in mechanosensitive channel gating following mechanical stimulation in skeletal muscle myotubes from the <i>mdx</i> mouse," <i>J Physiol.</i> , 2002;539.2:391-407.
	X	Geiger and Bershadsky, "Assembly and mechanosensory function and focal contacts," <i>Curr Opin Cell Biol.</i> , 2001;13:584-592.
	X	Greenberg et al., "Prostate Cancer in a Transgenic Mouse," <i>Proc Natl Acad Sci USA</i> , April 1995;92:3439-3443.

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	X	Grimes et al., "Differential expression of voltage-activated Na <sup>+</sup> currents in two prostatic tumor cell lines: contribution to invasiveness in vitro," <i>FEBS Letts.</i> , 1995;369:290-294.
	X	Guharay and Sachs, "Stretch-activated single ion channel currents in tissue cultured embryonic chick skeletal muscle," <i>J Physiol.</i> , 1984;352:685-701.
	X	Gutierrez et al., "Activation of a Ca <sup>2+</sup> -permeable cation channel by two different inducers of apoptosis in human prostatic cancer cell line," <i>J Physiol.</i> , May 1999;517.1:95-107.
	X	Haas & Sakr., "Epidemiology of Prostate Cancer," <i>CA Cancer Journal Clinic</i> , September/October 1997;47(5):273-287.
	X	Hamajima et al., "Intranasal administration of HIV-DNA vaccine formulated with a polymer, carboxymethylcellulose, augments mucosal antibody production and cell-mediated immune response," <i>Clin Immunol. Immunopathol.</i> , August 1998;88(2):205-210.
	X	Hamill & McBride, "Rapid adaptation of single mechanosensitive channels in <i>Xenopus</i> oocytes," <i>Proc Natl Acad Sci USA</i> , 15 August 1992;89:7462-7466.
	X	Hamill & McBride, "The Pharmacology of Mechanogated Membrane Ion Channels," <i>Pharmacological Reviews</i> , 1996;48(2):231-252.
	X	Hamill & McBride, "Induced membrane hypo/hyper-mechanosensitivity: a limitation of patch-clamp recording," <i>Ann Rev Physiol.</i> , 1997;59:621-631.
	X	Hamill & McBride, "Patch and whole cell currents recorded from BC3H-1 muscle cells," <i>Biophys J.</i> , February 14-18 1993; 37 <sup>th</sup> Annual Meeting Program and Abstracts, Washington D.C.;64:M-Pos 407;A93.
	X	Hamill & McBride, "Pressure/patch-clamp methods," <i>Patch Clamp Applications and Protocols</i> , Boulton eds., Humana Press Inc., New Jersey, 1995:75-87.

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	X	Hamill & Martinac, "Molecular basis of mechanotransduction in living cells," <i>Physiol Revs.</i> , April 2001;81:685-740.
	X	Hamill et al., "Improved patch-clamp techniques for high-resolution current recording from cells and cell-free membrane patches," <i>Pfluegers Archiv</i> , 1981;391:85-100.
	X	Hamill et al., "Calcium-activated apoptosis in frog ( <i>Xenopus laevis</i> ) red blood cells," <i>J Physiol.</i> , September 2000;527P:45-46P.
	X	Harteneck et al., "From worm to man: three subfamilies of TRP channels," <i>Trends Neurosci.</i> , 2000;23:159-166.
	X	Hofmann et al., "Subunit composition of mammalian transient receptor potential channels in living cells," <i>Proc Natl Acad Sci USA</i> , 28 May 2002;99(11):7461-7466.
	X	Hu et al., "Stretch-activated Ion Channels in the Heart," <i>J Mol Cell Cardiol.</i> , 1997;29:1511-1523.
	X	Huang et al., "Identification of Channels Promoting Calcium Spikes and Waves in HT1080 Tumor Cells: Their Apparent Roles in Cell Motility and Invasion," <i>Cancer Res</i> , 1 April 2004;64:2482-2489.
	X	Jacque et al., "Modulation of HIV-1 replication by RNA interference," <i>Nature</i> , 25 July 2002;418:435-438.
	X	Jespersen et al., "Dual function vector for protein expression in both mammalian cells and <i>Xenopus laevis</i> oocytes," <i>Biotechniques</i> , March 2002;32(3):536-540.
	X	Kassis et al., "Tumor invasion as dysregulated cell motility," <i>Cancer Biology</i> , 2001;11:105-117.
	X	Lane et al., "Structure-activity relations of amiloride and its analogues in blocking the mechanosensitive channel in <i>Xenopus</i> oocytes," <i>Brit J. Pharmacol.</i> , 1992;106(2)283-286.

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	X	Lane et al., "Amiloride block of the mechanosensitive cation channel in <i>Xenopus</i> oocytes," <i>J Physiol</i> , 1991;441:347-366.
	X	Lauffenburger & Horwitz, "Cell Migration: a physically integrated molecular process," <i>Cell</i> , 9 February 1996;84:359-369.
	X	Lee et al., "Regulation of cell movement is mediated by stretch-activated calcium channels," <i>Nature</i> , 22 July 1999;400:382-386.
	X	Lintschinger et al., "Coassembly of Trp1 and Trp3 Proteins Generates Diacylglycerol- and Ca <sup>2+</sup> -sensitive Cation Channels," <i>J Biol Chem</i> , 8 September 2000;275(36):27799-27805.
	X	Liotta, "Tumor Invasion and Metastases- Role of the Extracellular Matrix: Rhoads Memorial Award Lecture," <i>Cancer Research</i> , January 1986;46:1-7.
	X	Lockwich et al., "Assembly of Trp1 in a Signaling Complex Associated with Caveolin-Scaffolding Lipid Raft Domains," <i>J Biol Chem</i> , 21 April 2000;275(16):11934-11942.
	X	Mandeville et al., "Intracellular calcium levels correlate with speed and persistent forward motion in migrating neutrophils," <i>Biophys. J</i> , April 1995;68:1207-1217.
	X	Marks et al., "Transient Increases in Cytosolic Free Calcium Appear to be Required for the Migration of Adherent Human Neutrophils," <i>J Cell Biol</i> , 1990;110:43-52.
	X	Maroto and Hamill, "Brefeldin A block of Integrin-dependent Mechanosensitive ATP release from <i>Xenopus</i> Oocytes Reveals a Novel Mechanism of Mechanotransduction," <i>J Biol Chem</i> , 29 June 2001;276(26):23867-23872.
	X	Maroto et al., TRPC1 forms the stretch-activated cation channel in vertebrate cells," <i>Nature Cell Biol.</i> , February 2005;7(2):179-185.

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	X	Martinac et al., "Mechanosensitive ion channels of E. coli activated by amphipaths," <i>Nature</i> , 15 November 1990;348:261-263.
	X	Martinac & Hamill, "Gramicidin A channels switch between stretch activation and stretch inactivation depending upon bilayer thickness," <i>Proc. Natl. Acad. Sci. USA.</i> , 2 April 2002;99(7):4308-4312.
	X	Martinac & Kloda, "Evolutionary origins of mechanosensitive ion channels," <i>Prog. Biophys. Molec. Biol.</i> , 2003;82:11-24.
	X	McBride et al., "Pressure-clamp: a method for rapid step perturbation of mechanosensitive channels," <i>Pflugers Archiv.</i> , 1992;421:606-612.
	X	McBride and Hamill, "A fast pressure clamp technique for studying mechanogated channels," <i>Single Channel Recording</i> , Chapter 14, 2 <sup>nd</sup> Edition, Sakamann eds., Plenum Press, New York, NY, 1995:329-340.
	X	Mercurio et al., "Towards a mechanistic understanding of tumor invasion—lessons from the $\alpha 6 \beta 4$ integrin," <i>Semin. Cancer Biol.</i> , April 2001;11(2):129-141.
	X	Minke & Cook, "TRP chanel proteins and signal transduction," <i>Physiol. Rev.</i> , 2002;82:429-472.
	X	Montell C., "New light on TRP and TRPL," <i>Mol. Pharmacol.</i> , 1997;52:755-763.
	X	Munevar et al., "Regulation of mechanical interactions between fibroblasts and the substratum by stretch-activated $\text{Ca}^{2+}$ entry," <i>J Cell Science</i> , 2003;117:85-92.
	X	Narasimhan et al., "Snail and Spider toxins share a similar tertiary structure and 'cristine motif'," <i>Nature Structural Biol.</i> , 1994;1:850-852.

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	X	National Center for Biotechnology Information, National Library of Medicine, National Institutes of Health, GenBank Locus X89066, Accession No. X89066, "H. sapiens mRNA for TRPC1 protein," [online]. Bethesda, MD [retrieved on 2006-11-16]. Retrieved from the Internet:<URL:http://www.ncbi.nlm.nih.gov/entrez/viewer.fcgi?db=nucleotide&val=1370118>; 5 pgs.
	X	National Center for Biotechnology Information, National Library of Medicine, National Institutes of Health, GenBank Locus P48995, Accession No. P48995, "Short transient receptor potential channel 1 (TrpC1) (TRP-1 protein)," [online]. Bethesda, MD [retrieved on 2006-11-16]. Retrieved from the Internet:<URL:http://www.ncbi.nlm.nih.gov/entrez/viewer.fcgi?db=protein&val=1351302>; 6 pgs.
	X	National Center for Biotechnology Information, National Library of Medicine, National Institutes of Health, GenBank Locus CAA61447, Accession No. CAA61447, "TRPC1 Protein (Homo sapiens)," [online]. Bethesda, MD [retrieved on 2006-11-16]. Retrieved from the Internet:<URL:http://www.ncbi.nlm.nih.gov/entrez/viewer.fcgi?db=protein&val=1370119>;3pgs.
	X	National Center for Biotechnology Information, National Library of Medicine, National Institutes of Health, GenBank Locus A59371, Accession No. A59371, "Toxin GsMTx-4 (validated)- Chilean tarantula," [online]. Bethesda, MD [retrieved on 2006-11-16]. Retrieved from the Internet:<URL:http://www.ncbi.nlm.nih.gov/entrez/viewer.fcgi?db=protein&val=25412346>;2 pgs.
	X	Norton et al., "The cystine knot structure of ion channel toxins and related polypeptides," <i>Toxicon</i> , November 1998;36(11):1573-1583.
	X	Ostrow et al., "cDNA sequence and in vitro folding of GsMTx4, a specific peptide inhibitor of mechanosensitive channels," <i>Toxicon</i> , 2003;42:263-274.

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	X	Paria et al., "Tumor Necrosis Factor- $\alpha$ Induces Nuclear Factor- $\kappa$ B-dependent TRPC1 Expression in Endothelial Cells," <i>J Biol Chem</i> , 26 September 2003;278(39):37195-37203.
	X	Polin et al., "Treatment of human prostate tumors PC-3 and TSU-PR1 with standard and investigational agents in SCID mice," <i>Invest. New Drugs</i> , 1997;15(2):99-108.
	X	Rafelski et al., "Crawling Toward a unified model of Cell Motility: Spatial and Temporal Regulation of Actin Dynamics," <i>Annual Review Biochemistry</i> , 2004;73:209-239.
	X	Riccio et al., "mRNA distribution analysis of human TRPC family in CNS and peripheral tissues," <i>Molec. Brain Res.</i> , 2002;109:95-104.
	X	Ridley et al., "Cell Migration: Integrating signals from Front to Back," <i>Science</i> , 5 December 2003;302(5651):1704-1709.
	X	Sachs & Morris, "Mechanosensitive ion channels in nonspecialized cells," <i>Revs Physiol. Biochem &amp; Pharm.</i> , 1998;132:1-77.
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	<b>Applicant(s):</b> HAMILL et al.	<b>Confirmation No.:</b> 2236
	<b>Application Filing Date:</b> July 7, 2006	<b>Group:</b> 1642
	<b>Information Disclosure Statement mailed:</b> November 22, 2006	

Examiner Initial	Copy Enclosed	Document Description
	X	Smith et al., "Sodium channel protein expression enhances the invasiveness of rat and human prostate cancer cells," <i>FEBS Letts.</i> , 1998;423:19-24.
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